1040 LQFLGFWGNRVGDEGAQALAEALGDHQSLRW.SLVGNNI GSVGAQALALMAKNVM.EELCLEENHLQDEGVCSLAEGLKKNSSLKI LKLSNNGTYLGA 1000 **QHVQELPVPLALPLEAATCK**KYNAKLRTTVSAQSRFLSTYDGAETLCLEDI YTENVLEVWADVGNAGPPQKSPATLGLEELFSTPGHLNDDADTVLVV任 300 AGSGKS]TLLQRLHLLWAAQQXQEPLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRV<u>LLTFD</u>GFDEFKFRFTDRERHCSPTDP 400 TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYI RTEFNLKGFSEQGI ELYLRKRHHEPGVADRLI RLLQETSALHGLCHLPVFSWM/SKCHQELL 500 LQEGGSPKITIDMALLI LQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGMCCYVFSAQQLQAAQVSPDDI SLGFLVRAKGVVPGSTAPLE 600 FLHI TFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPMARLLPTMCI QASEGKDSSVAALLQKAEPHNLQI TAAFLAGLLSREHWGLLAECQTSEKAL 700 LRHQACARWCLARSLRKHFHSI PPAAPGEAKSVHAMPGFI W.I RSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN 800 SVGDI GVEQLLPCLGVCKALYLRDNNI SDRGI CKLI ECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGNNYI TAAGAQVLAEGLRGNTS 900 QKLI AAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAI VRRLHSHVENMLDLAWERGFVSQYECDEI RLPI FTPSQRARRLLDLATVKANGLAAFLL 200 MÆEGGS AS HDEEBRAS VLLCHSP GCE<mark>MCS QEAF QAQRS QLVELLVS GSLEGFES VLDWLLS WEVLS WEDYEGFHLLGQPLS HLARRLLDTVWNKGTWAC</mark> 100 EALLOÁLERNDTI LEVWLRGNTFSLEEVDKLGCRDTRLLL

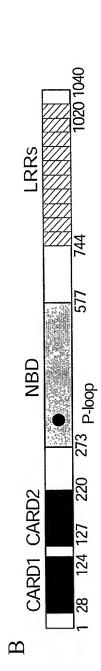
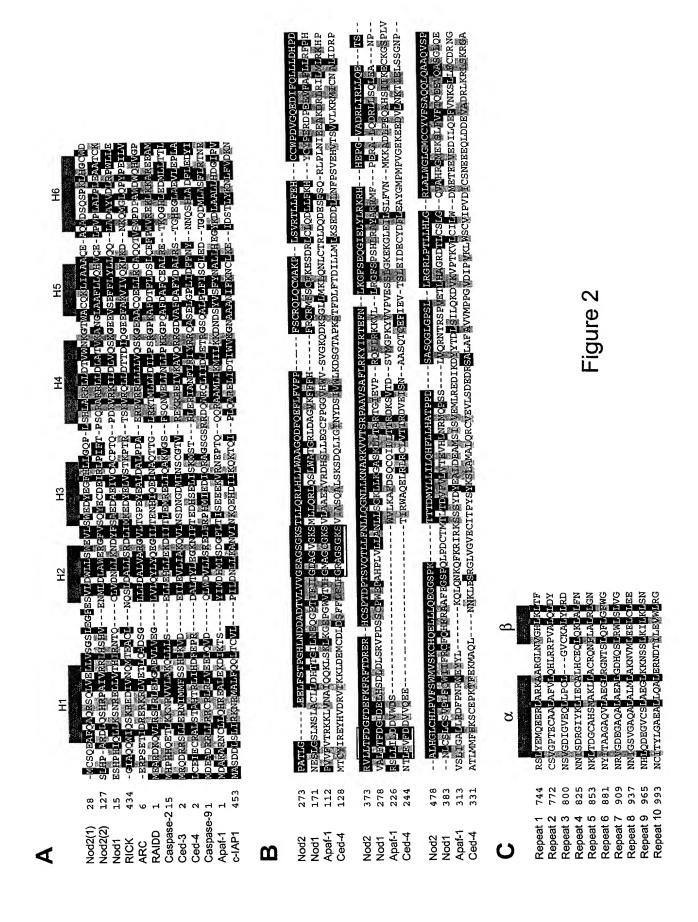


Figure 1



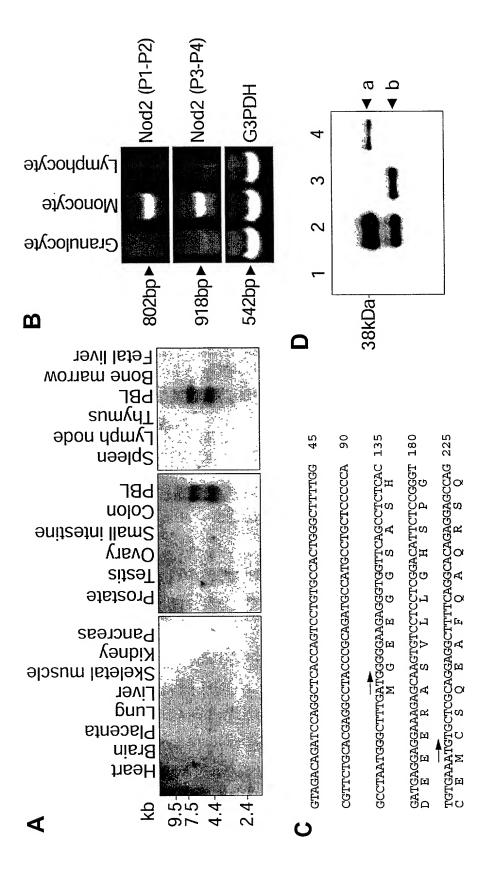


Figure 3

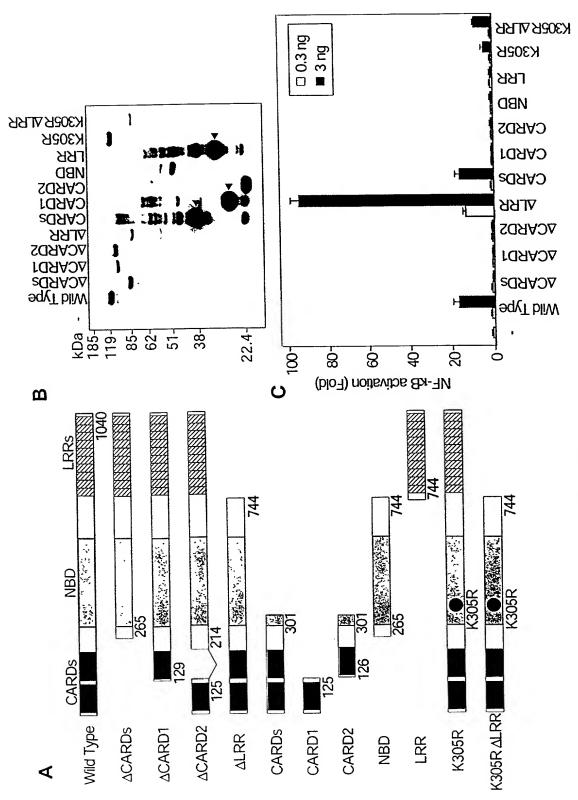


Figure 4

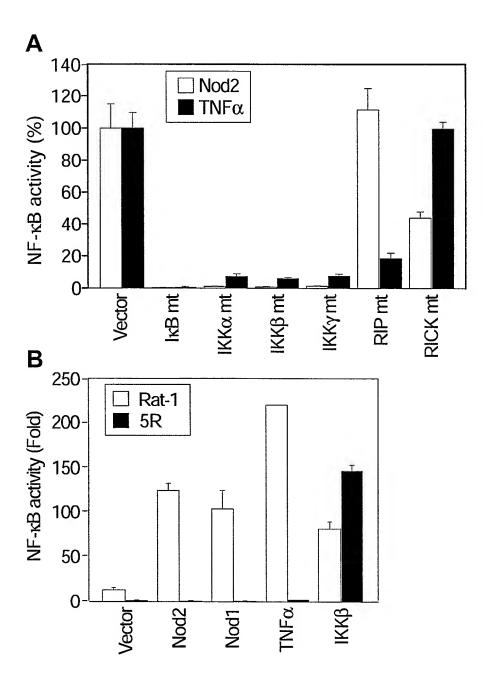
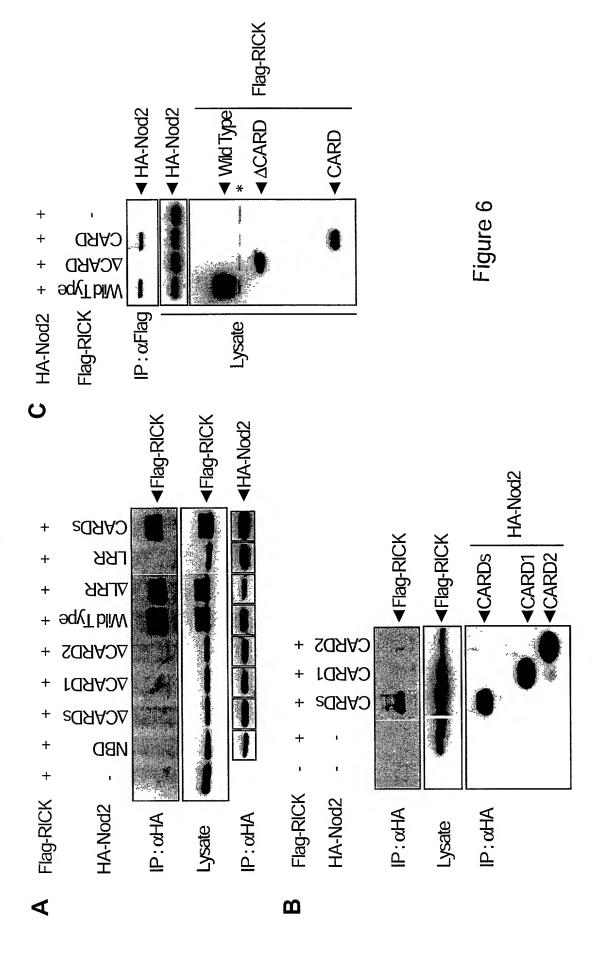


Figure 5



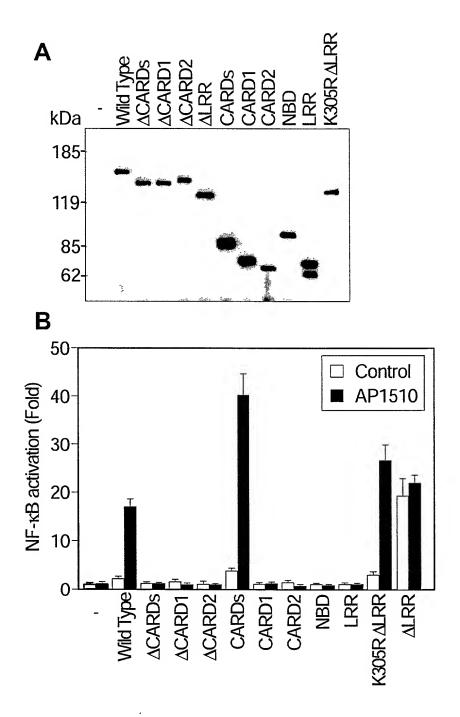


Figure 7

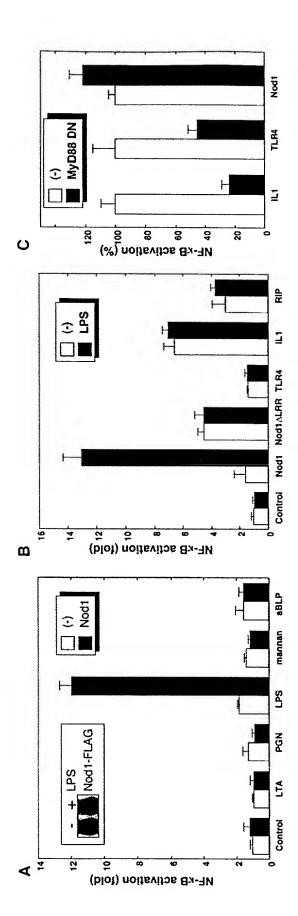


Figure 8

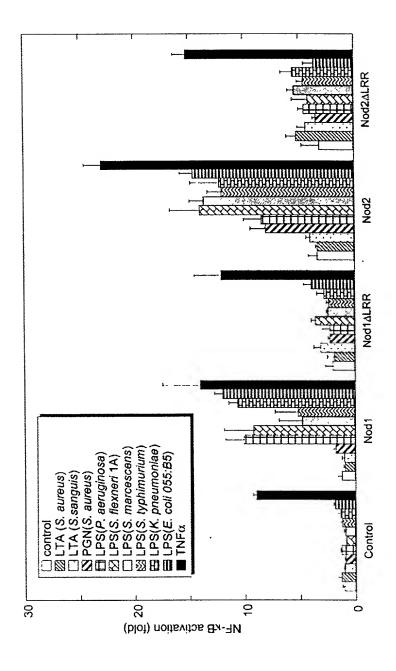


Figure 9

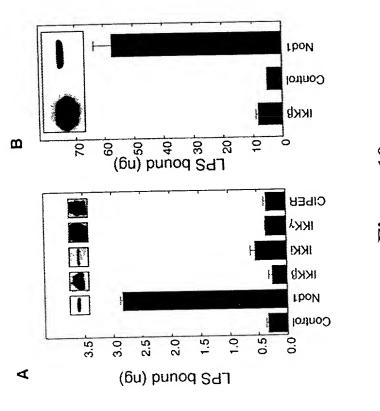


Figure 10

## Figure 11

### **SEQ ID NO:33**

#### Nod2 cDNA sequence

gtagacagat	ccaggctcac	cagtcctgtg	ccactgggct	tttggcgttc	tgcacaaggc	60
ctacccgcag	atgccatgcc	tgctccccca	gcctaatggg	ctttgatggg	ggaagagggt	120
ggttcagcct	ctcacgatga	ggaggaaaga	gcaagtgtcc	tcctcggaca	tteteegggt	180
tgtgaaatgt	gctcgcagga	ggcttttcag	gcacagagga	gccagctggt	cgagctgctg	240
gtctcagggt	ccctggaagg	cttcgagagt	gtcctggact	ggctgctgtc	ctgggaggtc	300
ctctcctggg	aggactacga	gggcttccac	ctcctgggcc	agcctctctc	ccacttggcc	360
aggegeette	tggacaccgt	ctggaataag	ggtacttggg	cctgtcagaa	gctcatcgcg	420
gctgcccaag	aagcccaggc	cgacagccag	tcccccaagc	tgcatggctg	ctgggacccc	480
cactcgctcc	acccagcccg	agacctgcag	agtcaccggc	cagccattgt	caggaggctc	540
cacagccatg	tggagaacat	gctggacctg	gcatgggagc	ggggtttcgt	cagccagtat	600
gaatgtgatg	aaatcaggtt	gccgatcttc	acaccgtccc	agagggcaag	aaggctgctt	660
gatcttgcca	cggtgaaagc	gaatggattg	getgeettee	ttctacaaca	tgttcaggaa	720
ttaccagtcc	cattggccct	gcctttggaa	gctgccacat	gcaagaagta	tatggccaag	780
ctgaggacca	cggtgtctgc	tcagtctcgc	ttcctcagta	cctatgatgg	agcagagacg	840
ctctgcctgg	aggacatata	cacagagaat	gtcctggagg	tctgggcaga	tgtgggcatg	900
gctggacccc	cgcagaagag	cccagccacc	: ctgggcctgg	aggagctctt	cagcacccct	960
ggccacctca	. atgacgatgo	ggacactgtg	ctggtggtgg	gtgaggcggg	cagtggcaag	1020
agcacgctcc	tgcagcggct	gcacttgctg	ı tgggctgcag	ggcaagactt	ccaggaattt	1080
ctctttgtct	tcccattcag	r ctgccggcag	g ctgcagtgca	tggccaaacc	actctctgtg	1140
cggactctac	tetttgagea	ctgctgttgg	cctgatgttg	gtcaagaaga	a catcttccag	1200
ttactccttg	g accaccctga	ccgtgtcctg	g ttaacctttg	g atggetttga	a cgagttcaag	1260
ttcaggttca	a cqqatcqtqa	acqccactq	tccccgacco	accccaccto	tgtccagacc	1320

1380 ctgctcttca accttctgca gggcaacctg ctgaagaatg cccgcaaggt ggtgaccagc cgtccggccg ctgtgtcggc gttcctcagg aagtacatcc gcaccgagtt caacctcaag 1440 ggottototg aacagggcat cgagotgtac otgaggaago gocatcatga gocoggggtg 1500 1560 geggaeegee teateegeet getecaagag aceteageee tgeaeggttt gtgeeacetg cctgtcttct catggatggt gtccaaatgc caccaggaac tgttgctgca ggagggggg 1620 tececaaaga eeactacaga tatgtaeetg etgattetge ageattttet getgeatgee 1680 1740 accececag acteagette ecaaggtetg ggacceagte ttettegggg ecgeeteece accetectge acctgggeag actggetetg tggggeetgg geatgtgetg etacgtgtte 1800 1860 teageceage agetecagge ageacaggte agecetgatg acatttetet tggetteetg gtgcgtgcca aaggtgtcgt gccagggagt acggcgcccc tggaattcct tcacatcact 1920 1980 ttccagtget tetttgeege gttetacetg geacteagtg etgatgtgee accagetttg 2040 ctcagacacc tottcaattg tggcaggcca ggcaactcac caatggccag getectgccc 2100 acgatgtgca tccaggcctc ggagggaaag gacagcagcg tggcagcttt gctgcagaag 2160 gccgagccgc acaacettca gatcacagca gccttcctgg cagggctgtt gtcccgggag cactggggcc tgctggctga gtgccagaca tctgagaagg ccctgctccg gcgccaggcc 2220 2280 tgtgcccgct ggtgtctggc ccgcagcctc cgcaagcact tccactccat cccgccagct gcaccgggtg aggccaagag cgtgcatgcc atgcccgggt tcatctggct catccggagc 2340 2400 ctgtacgaga tgcaggagga gcggctggct cggaaggctg cacgtggcct gaatgttggg cacctcaagt tgacattttg cagtgtgggc cccactgagt gtgctgccct ggcctttgtg 2460 ctgcagcacc tccggcggcc cgtggccctg cagctggact acaactctgt gggtgacatt 2520 2580 ggcgtggagc agctgctgcc ttgccttggt gtctgcaagg ctctgtattt gcgcgataac 2640 aatateteaq aeegaggeat etgeaagete attgaatgtg etetteaetg egageaattg 2700 cagaagttag ctctattcaa caacaaattg actgacggct gtgcacactc catggctaag ctccttgcat gcaggcagaa cttcttggca ttgaggctgg ggaataacta catcactgcc 2760 2820 gegggagece aagtgetgge egaggggete egaggeaaca ceteettgea gtteetggga 2880 ttctggggca acagagtggg tgacgagggg gcccaggccc tggctgaagc cttgggtgat 2940 caccagaget tgaggtgget cageetggtg gggaacaaca ttggcagtgt gggtgeecaa 3000 gccttggcac tgatgctggc aaagaacgtc atgctagaag aactctgcct ggaggagaac 3060 catctccagg atgaaggtgt atgttctctc gcagaaggac tgaagaaaaa ttcaagtttg 3120 aaaatcctga agttgtccaa taactgcatc acctacctag gggcagaagc cctcctgcag

3180 gccccttgaaa ggaatgacac catcctggaa gtctggctcc gagggaacac tttctctcta gaggaggttg acaagctcgg ctgcagggac accagactct tgctttgaag tctccgggag 3240 3300 gatgttcgtc tcagtttgtt tgtgagcagg ctgtgagttt gggccccaga ggctgggtga catgtgttgg cagcetette aaaatgagee etgteetgee taaggetgaa ettgttteet 3360 gggaacacca taggtcacct ttattctggc agaggaggga gcatcagtgc cctccaggat 3420 3480 agacttttcc caagcctact tttgccattg acttcttccc aagattcaat cccaggatgt acaaggacag eccetectee atagtatggg actggeetet getgateete ecaggettee 3540 3600 gtgtgggtca gtggggccca tggatgtgct tgttaactga gtgccttttg gtggagaggc 3660 ccggcctctc acaaaagacc ccttaccact gctctgatga agaggagtac acagaacaca 3720 taattcagga agcagettte eccatgtete gaeteateea tecaggeeat teecegtete 3780 tggttcctcc cctcctcctg gactcctgca cacgctcctt cctctgaggc tgaaattcag aatattagtg acctcagett tgatatttca ettacageae eeccaaceet ggeacecagg 3840 gtgggaaggg ctacacctta gcctgccctc ctttccggtg tttaagacat ttttggaagg 3900 ggacacgtga cagccgtttg ttccccaaga cattctaggt ttgcaagaaa aatatgacca 3960 4020 cactccagct gggatcacat gtggactttt atttccagtg aaatcagtta ctcttcagtt aagcetttgg aaacageteg aetttaaaaa geteeaaatg cagetttaaa aaattaatet 4080 gggccagaat ttcaaacggc ctcactaggc ttctggttga tgcctgtgaa ctgaactctg 4140 acaacagact totgaaatag accoacaaga ggcagttoca tttcatttgt gccagaatgc 4200 tttaggatgt acagttatgg attgaaagtt tacaggaaaa aaaattaggc cgttccttca 4260 4320 aagcaaatgt cttcctggat tattcaaaat gatgtatgtt gaagcctttg taaattgtca gatgctgtgc aaatgttatt attttaaaca ttatgatgtg tgaaaactgg ttaatattta 4380 4440 taggtcactt tgttttactg tcttaagttt atactcttat agacaacatg gccgtgaact ttatgctgta aataatcaga ggggaataaa ctgttgagtc aaaac 4485

## Figure 12

## **SEQ ID NO:1**

#### Nod2 cDNA sequence

gta <u>g</u> aca	gat	ccaggctcac	cagtcctgtg	ccactgggct	tttggcgttc	tgcacaaggc	60
ctacccg	cag	atgccatgcc	tgctccccca	gcctaatggg	ctttgatggg	ggaagagggt	120
ggttcag	cct	ctcacgatga	ggaggaaaga	gcaagtgtcc	tcctcggaca	ttctccgggt	180
tgtgaaa	tgt	gctcgcagga	ggcttttcag	gcacagagga	gccagctggt	cgagctgctg	240
gtctcag	ggt	ccctggaagg	cttcgagagt	gtcctggact	ggctgctgtc	ctgggaggtc	300
ctctcct	999	aggactacga	gggcttccac	ctcctgggcc	agcctctctc	ccacttggcc	360
aggcgcc	ttc	tggacaccgt	ctggaataag	ggtacttggg	cctgtcagaa	gctcatcgcg	420
gctgccc	aag	aagcccaggc	cgacagccag	tececcaage	tgcatggctg	ctgggacccc	480
cactego	tcc	acccagcccg	agacctgcag	agtcaccggc	cagccattgt	caggaggctc	540
cacagco	atg	tggagaacat	gctggacctg	gcatgggagc	ggggtttcgt	cagccagtat	600
gaatgt	gatg	aaatcaggtt	gccgatcttc	acaccgtccc	agagggcaag	aaggctgctt	660
gatctt	gcca	cggtgaaagc	gaatggattg	gctgccttcc	ttctacaaca	tgttcaggaa	720
ttacca	gtcc	cattggccct	gcctttggaa	gctgccacat	gcaagaagta	tatggccaag	780
ctgagg	acca	cggtgtctgc	tcagtctcgc	: ttcctcagta	cctatgatgg	g agcagagacg	840
ctctgc	ctgg	aggacatata	ı cacagagaat	gtcctggagg	tctgggcaga	tgtgggcatg	900
gctgga	cccc	cgcagaagag	cccagccacc	ctgggcctgg	aggagctctt	cagcacccct	960
ggccac	ctca	. atgacgatgo	ggacactgtg	g ctggtggtgg	gtgaggcgg	g cagtggcaag	102
agcacg	ctcc	: tgcagcggct	gcacttgcto	g tgggctgcag	g ggcaagacti	ccaggaattt	108
ctcttt	gtct	teccattea	g ctgccggcag	g ctgcagtgca	a tggccaaac	c actetetgtg	114
cggact	ctac	totttgago	a ctgctgttg	g cctgatgtts	g gtcaagaag	a catcttccag	120
ttactc	ctt	g accaccctg	a ccgtgtcct	g ttaaccttt	g atggctttg	a cgagttcaag	126
ttcagg	ttca	cggatcgtg	a acgccactg	c teceegace	g accccacct	c tgtccagacc	132

1380 ctgctcttca accttctgca gggcaacctg ctgaagaatg cccgcaaggt ggtgaccagc 1440 cgtccggccg ctgtgtcggc gttcctcagg aagtacatcc gcaccgagtt caacctcaag ggettetetg aacagggeat egagetgtae etgaggaage gecateatga geceggggtg 1500 geggacegee teateegeet getecaagag aceteageee tgeaeggttt gtgeeaeetg 1560 cctgtcttct catggatggt gtccaaatgc caccaggaac tgttgctgca ggagggggg 1620 1680 tocccaaaga ccactacaga tatgtacctg ctgattctgc agcattttct gctgcatgcc accccccag actcagette ccaaggtetg ggacccagte ttettegggg eegecteece 1740 accetectge acctgggeag actggetetg tggggeetgg geatgtgetg etacgtgtte 1800 1860 tragerrage agetreage agearages agerrages acatterer tegetteres gtgcgtgcca aaggtgtcgt gccagggagt acggcgcccc tggaattcct tcacatcact 1920 1980 ttccagtgct tctttgccgc gttctacctg gcactcagtg ctgatgtgcc accagctttg 2040 ctcagacacc tcttcaattg tggcaggcca ggcaactcac caatggccag gctcctgccc acgatgtgca tccaggcctc ggagggaaag gacagcagcg tggcagcttt gctgcagaag 2100 2160 gccgagccgc acaacettca gatcacagca gccttcctgg cagggctgtt gtcccgggag 2220 cactggggcc tgctggctga gtgccagaca tctgagaagg ccctgctccg gcgccaggcc 2280 tgtgcccgct ggtgtctggc ccgcagcctc cgcaagcact tccactccat cccgccagct gcaccgggtg aggccaagag cgtgcatgcc atgcccgggt tcatctggct catccggagc 2340 2400 ctgtacgaga tgcaggagga gcggctggct cggaaggctg cacgtggcct gaatgttggg cacctcaagt tgacattttg cagtgtgggc cccactgagt gtgctgccct ggcctttgtg 2460 2520 ctgcagcacc tccggcggcc cgtggccctg cagctggact acaactctgt gggtgacatt ggcgtggagc agctgctgcc ttgccttggt gtctgcaagg ctctgtattt gcgcgataac 2580 aatatctcag accgaggcat ctgcaagctc attgaatgtg ctcttcactg cgagcaattg 2640 2700 cagaagttag ctctattcaa caacaaattg actgacggct gtgcacactc catggctaag ctccttgcat gcaggcagaa cttcttggca ttgaggctgg ggaataacta catcactgcc 2760 2820 gegggageee aagtgetgge egagggete egaggeaaca eeteettgea gtteetggga ttctggggca acagagtggg tgacgagggg gcccaggccc tggctgaagc cttgggtgat 2880 caccagaget tgaggtgget cagectggtg gggaacaaca ttggcagtgt gggtgeecaa 2940 3000 gccttggcac tgatgctggc aaagaacgtc atgctagaag aactctgcct ggaggagaac catctccagg atgaaggtgt atgttctctc gcagaaggac tgaagaaaaa ttcaagtttg 3060 3120 aaaatcctga agttgtccaa taactgcatc acctacctag gggcagaagc cctcctgcag

3180 gecettgaaa ggaatgacae cateetggaa gtetggetee gagggaacae ttteteteta gaggaggttg acaagctcgg ctgcagggac accagactct tgctttgaag tctccgggag 3240 3300 gatgttegte teagtttgtt tgtgageagg etgtgagttt gggeeceaga ggetgggtga catgtgttgg cagcctcttc aaaatgagcc ctgtcctgcc taaggctgaa cttgttttct 3360 3420 gggaacacca taggtcacct ttattctggc agaggaggga gcatcagtgc cctccaggat agacttttcc caagcctact tttgccattg acttcttccc aagattcaat cccaggatgt 3480 3540 acaaggacag cccctcctcc atagtatggg actggcctct gctgatcctc ccaggcttcc gtgtgggtca gtggggccca tggatgtgct tgttaactga gtgccttttg gtggagaggc 3600 3660 ccggcctctc acaaaagacc ccttaccact gctctgatga agaggagtac acagaacaca 3720 taattcagga agcagctttc cccatgtctc gactcatcca tccaggccat tccccgtctc 3780 tggttcctcc cctcctcctg gactcctgca cacgctcctt cctctgaggc tgaaattcag aatattagtg acctcagctt tgatatttca cttacagcac ccccaaccct ggcacccagg 3840 gtgggaaggg ctacacctta gcctgccctc ctttccggtg tttaagacat ttttggaagg 3900 3960 ggacacgtga cagccgtttg ttccccaaga cattctaggt ttgcaagaaa aatatgacca 4020 cactccaget gggatcacat gtggaetttt atttccagtg aaatcagtta etettcagtt 4080 aagcetttgg aaacageteg aetttaaaaa getecaaatg cagetttaaa aaattaatet 4140 gggccagaat ttcaaacggc ctcactaggc ttctggttga tgcctgtgaa ctgaactctg acaacagact tetgaaatag acceacaaga ggeagtteea ttteatttgt geeagaatge 4200 4260 tttaggatgt acagttatgg attgaaagtt tacaggaaaa aaaattaggc cgttccttca aagcaaatgt cttcctggat tattcaaaat gatgtatgtt gaagcctttg taaattgtca 4320 4380 gatgctgtgc aaatgttatt attttaaaca ttatgatgtg tgaaaactgg ttaatattta 4440 taggtcactt tgttttactg tcttaagttt atactcttat agacaacatg gccgtgaact 4485 ttatgctgta aataatcaga ggggaataaa ctgttgagtc aaaac

## Figure 13

## SEQ ID NO:2

MGEEGGSASH	DEEERASVLL	GHSPGCEMCS	QEAFQAQRSQ	LVELLVSGSL	EGFESVLDWL
LSWEVLSWED	YEGFHLLGQP	LSHLARRLLD	TVWNKGTWAC	QKLIAAAQEA	QADSQSPKLH
GCWDPHSLHP	ARDLQSHRPA	IVRRLHSHVE	NMLDLAWERG	FVSQYECDEI	RLPIFTPSQR
ARRLLDLATV	KANGLAAFLL	QHVQELPVPL	ALPLEAATCK	KYMAKLRTTV	SAQSRFLSTY
DGAETLCLED	IYTENVLEVW	ADVGMAGPPQ	KSPATLGLEE	LFSTPGHLND	DADTVLVVGE
AGSGKSTLLQ	RLHLLWAAGQ	DFQEFLFVFP	FSCRQLQCMA	KPLSVRTLLF	EHCCWPDVGQ
EDIFQLLLDH	PDRVLLTFDG	FDEFKFRFTD	RERHCSPTDP	TSVQTLLFNL	LQGNLLKNAR
KVVTSRPAAV	SAFLRKYIRT	EFNLKGFSEQ	GIELYLRKRH	HEPGVADRLI	RLLQETSALH
GLCHLPVFSW	MVSKCHQELL	LQEGGSPKTT	TDMYLLILQH	FLLHATPPDS	ASQGLGPSLL
RGRLPTLLHL	GRLALWGLGM	CCYVFSAQQL	QAAQVSPDDI	SLGFLVRAKG	VVPGSTAPLE
FLHITFQCFF	AAFYLALSAD	VPPALLRHLF	NCGRPGNSPM	ARLLPTMCIQ	ASEGKDSSVA
ALLQKAEPHN	LQITAAFLAG	LLSREHWGLL	AECQTSEKAL	LRRQACARWC	LARSLRKHFH
SIPPAAPGEA	KSVHAMPGFI	WLIRSLYEMQ	EERLARKAAR	GLNVGHLKLT	FCSVGPTECA
ALAFVLQHLR	RPVALQLDYN	SVGDIGVEQL	LPCLGVCKAL	YLRDNNISDR	GICKLIECAL
HCEQLQKLAL	FNNKLTDGCA	HSMAKLLACR	QNFLALRLGN	NYITAAGAQV	LAEGLRGNTS
LQFLGFWGNR	VGDEGAQALA	EALGDHQSLR	WLSLVGNNIG	SVGAQALALM	LAKNVMLEEL
CLEENHLQDE	GVCSLAEGLK	KNSSLKILKL	SNNCITYLGA	EALLQALERN	DTILEVWLRG
NTFSLEEVDK	LGCRDTRLLL	*			

#### Figure 14

#### SEQ ID NO:3

MCSQEAFQAQ RSQLVELLVS GSLEGFESVL DWLLSWEVLS WEDYEGFHLL GQPLSHLARR LLDTVWNKGT WACQKLIAAA QEAQADSQSP KLHGCWDPHS LHPARDLQSH RPAIVRRLHS HVENMLDLAW ERGFVSQYEC DEIRLPIFTP SQRARRLLDL ATVKANGLAA FLLQHVQELP VPLALPLEAA TCKKYMAKLR TTVSAQSRFL STYDGAETLC LEDIYTENVL EVWADVGMAG PPQKSPATLG LEELFSTPGH LNDDADTVLV VGEAGSGKST LLQRLHLLWA AGQDFQEFLF VFPFSCRQLQ CMAKPLSVRT LLFEHCCWPD VGQEDIFQLL LDHPDRVLLT FDGFDEFKFR FTDRERHCSP TDPTSVQTLL FNLLQGNLLK NARKVVTSRP AAVSAFLRKY IRTEFNLKGF SEQGIELYLR KRHHEPGVAD RLIRLLQETS ALHGLCHLPV FSWMVSKCHQ ELLLQEGGSP KTTTDMYLLI LQHFLLHATP PDSASQGLGP SLLRGRLPTL LHLGRLALWG LGMCCYVFSA OOLOAAOVSP DDISLGFLVR AKGVVPGSTA PLEFLHITFQ CFFAAFYLAL SADVPPALLR HLFNCGRPGN SPMARLLPTM CIQASEGKDS SVAALLQKAE PHNLQITAAF LAGLLSREHW GLLAECQTSE KALLRRQACA RWCLARSLRK HFHSIPPAAP GEAKSVHAMP GFIWLIRSLY EMOEERLARK AARGLNVGHL KLTFCSVGPT ECAALAFVLQ HLRRPVALQL DYNSVGDIGV EQLLPCLGVC KALYLRDNNI SDRGICKLIE CALHCEQLQK LALFNNKLTD GCAHSMAKLL ACRONFLALR LGNNYITAAG AQVLAEGLRG NTSLQFLGFW GNRVGDEGAQ ALAEALGDHQ SLRWLSLVGN NIGSVGAQAL ALMLAKNVML EELCLEENHL QDEGVCSLAE GLKKNSSLKI LKLSNNCITY LGAEALLQAL ERNDTILEVW LRGNTFSLEE VDKLGCRDTR LLL\*

## Figure 15 SEQ ID NO:34

#### Nod2a AA sequence, Mutant

MGEEGGSASH DEEERASVLL GHSPGCEMCS QEAFQAQRSQ LVELLVSGSL EGFESVLDWL LSWEVLSWED YEGFHLLGQP LSHLARRLLD TVWNKGTWAC QKLIAAAQEA QADSQSPKLH GCWDPHSLHP ARDLQSHRPA IVRRLHSHVE NMLDLAWERG FVSQYECDEI RLPIFTPSQR ARRLLDLATV KANGLAAFLL QHVQELPVPL ALPLEAATCK KYMAKLRTTV SAQSRFLSTY DGAETLCLED IYTENVLEVW ADVGMAGPPQ KSPATLGLEE LFSTPGHLND DADTVLVVGE AGSGKSTLLQ RLHLLWAAGQ DFQEFLFVFP FSCRQLQCMA KPLSVRTLLF EHCCWPDVGQ EDIFQLLLDH PDRVLLTFDG FDEFKFRFTD RERHCSPTDP TSVQTLLFNL LQGNLLKNAR KVVTSRPAAV SAFLRKYIRT EFNLKGFSEQ GIELYLRKRH HEPGVADRLI RLLQETSALH GLCHLPVFSW MVSKCHQELL LQEGGSPKTT TDMYLLILQH FLLHATPPDS ASQGLGPSLL RGRLPTLLHL GRLALWGLGM CCYVFSAQQL QAAQVSPDDI SLGFLVRAKG VVPGSTAPLE FLHITFOCFF AAFYLALSAD VPPALLRHLF NCGRPGNSPM ARLLPTMCIQ ASEGKDSSVA ALLQKAEPHN LQITAAFLAG LLSREHWGLL AECQTSEKAL LRRQACARWC LARSLRKHFH SIPPAAPGEA KSVHAMPGFI WLIRSLYEMQ EERLARKAAR GLNVGHLKLT FCSVGPTECA ALAFVLQHLR RPVALQLDYN SVGDIGVEQL LPCLGVCKAL YLRDNNISDR GICKLIECAL HCEQLQKLAL FNNKLTDGCA HSMAKLLACR QNFLALRLGN NYITAAGAQV LAEGLRGNTS LQFLGFWGNR VGDEGAQALA EALGDHQSLR WLSLVGNNIG SVGAQALALM LAKNVMLEEL CLEENHLQDE GVCSLAEGLK KNSSLKILKL SNNCITYLGA EALLQAP\*

#### FIGURE 16

Nod2 Exon11, Wild type

cagacatgag caggatgtgt ctaagggaca ggtgggcttc agtagactgg ctaactcetg

cagtctcttt aactggacag tttcaagagg aaaaccaaga atcettgaag ctcaccattg

tatcttcttt tccagGTTGT CCAATAACTG CATCACCTAC CTAGGGGCAG AAGCCCTCCT

L S N N C I T Y L G A E A L L

GCAGGCCCTT GAAAGGAATG ACACCATCCT GGAAGTCTGg taaggccct gggcaggcct

Q A L E R N D T I L E V

gttttagctc tccgaacctc agtttttcta tctgtaaaat ggggtgacgg gagagagaa

tggcagaatt ttgaggatcc cttctgattc tgacattcag tgagaatgat tctgcatgtg

Nod2 Exon11, Mutant

cagacatgag caggatgtgt ctaagggaca ggtgggcttc agtagactgg ctaactcctg

cagtctcttt aactggacag tttcaagagg aaaaccaaga atccttgaag ctcaccattg

tatcttcttt tccaggttgt CCAATAACTG CATCACCTAC CTAGGGGCAG AAGCCCTCCT

L S N N C I T Y L G A E A L L

GCAGGCCCCT TGAAAGGAAT GACACCATCC TGGAAGTCTG gtaaggccc tgggcaggcc

#### Q A <u>P</u> <u>\*</u>

tgttttagct ctccgaacct cagtttttct atctgtaaaa tggggtgacg ggagagaga atggcagaat tttgaggatc ccttctgatt ctgacattca gtgagaatga ttctgcatgt

g

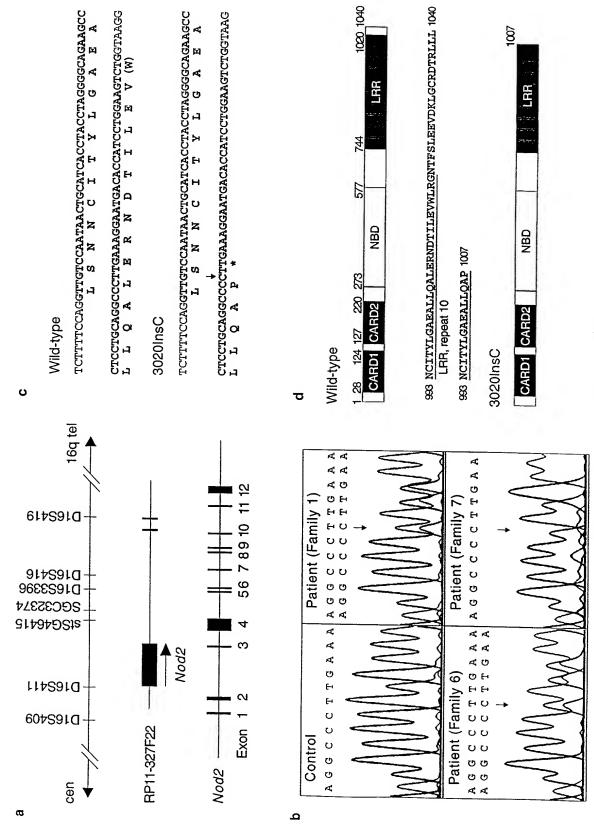


Figure 17

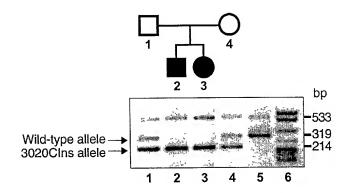


Figure 18

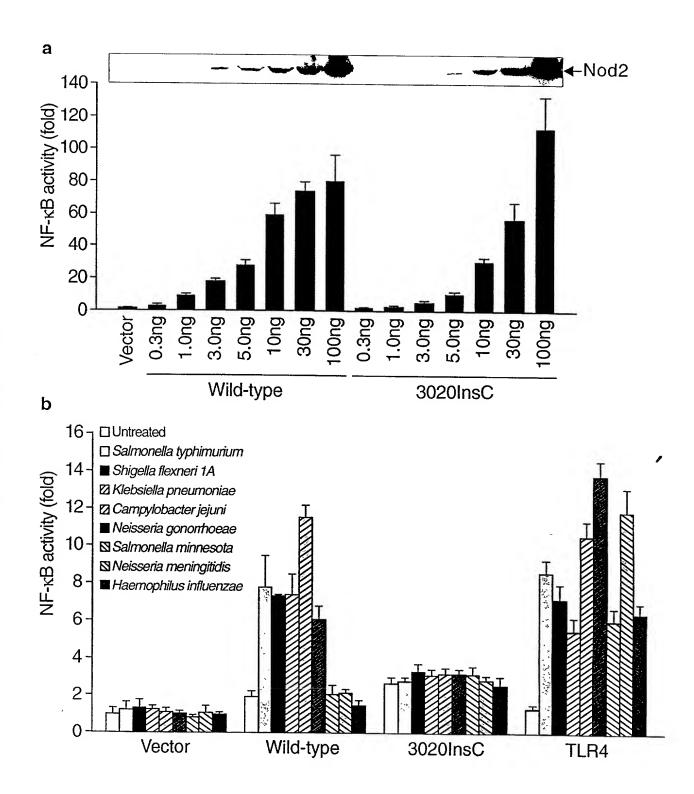


Figure 19

# Figure 20 SEQ ID NO: 53

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg agagtgtcctggactggctgctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc ccaagaagcccaggccgacagccagtccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg accccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgggtgaggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac  $\verb|tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg|$ acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg tgctgctacgtgttctcagcccagcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga agacatctgagaaggccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac tctgtgggtgacattggcgtggagcagctgctgccttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg gggcaacagagtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc  $\verb|tggtgggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc|$ tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgctt tga

# Figure 21 SEQ ID NO: 54

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga  $\verb| aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg| \\$ agagtqtcctqqactqqctqtcttgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct ctctcccacttqqccaqqcqccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc ccaagaagcccaggccgacagccagtccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tqccacqqtqaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg atccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgggtgaggggagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac  $\verb|tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg|$ acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccqtcc qqccqctqtqtcqqcqttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tqtacctqaqqaaqcqccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac qqtttqtqccacctqcctqtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggagggggggtcccc aaaqaccactacaqatatqtacctgctgattctgcagcatttttctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg tqctqctacqtqttctcaqcccaqcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg  ${\tt tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct}$ acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg qccaqqctcctqcccacqatqtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga  $\tt gccgcacaaccttcagatcacagcagccttcctggcagggctgttgtcccgggagcactggggcctgctgagtgcc$ agacatctgagaaggccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac tctgtgggtgacattggcgtggagcagctgctgccttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg gggcaacagagtgggtgacgagggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcaqtgtgggcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggccccttgaaaggaatgacaccatc  $\verb|ctggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgct|$ ttga

# Figure 22 SEQ ID NO: 55

		SH DEEERASVL	L GHSPGCEMCS	QEAFQAQRSQ	LVELLVSGSL	EGFESVLDWL
60		ED YEGFHLLGQ	P LSHLARRLLD	TVWNKGTWAC	QKLIAAAQEA	QADSQSPKLH
120		HP ARDLQSHRP	A IVRRLHSHVE	NMLDLAWERG	FVSQYECDEI	RLPIFTPSQR
180		TV KANGLAAFL	L QHVQELPVPL	ALPLEAATCK	KYMAKLRTTV	SAQSRFLSTY
240		ED IYTENVLEV	W ADVGMAG <u>s</u> PQ	KSPATLGLEE	LFSTPGHLND	DADTVLVVGE
300		LQ RLHLLWAAG	Q DFQEFLFVFP	FSCRQLQCMA	KPLSVRTLLF	EHCCWPDVGQ
360		DH PDRVLLTFD	G FDEFKFRFTD	RERHCSPTDP	TSVQTLLFNL	LQGNLLKNAR
420		AV SAFLRKYIR	T EFNLKGFSEQ	GIELYLRKRH	HEPGVADRLI	RLLQETSALH
480	GLCHLPVF	'SW MVSKCHQEL	L LQEGGSPKTT	' TDMYLLILQH	FLLHATPPDS	ASQGLGPSLL
540		HL GRLALWGLG	M CCYVFSAQQI	QAAQVSPDDI	SLGFLVRAKG	VVPGSTAPLE
600		CFF AAFYLALSA	D VPPALLRHLF	' NCGRPGNSPM	ARLLPTMCIQ	ASEGKDSSVA
661		PHN LQITAAFLA	G LLSREHWGLI	AECQTSEKAL	LRRQACARWC	LARSLRKHFH
72		SEA KSVHAMPGF	'I WLIRSLYEMÇ	) EERLARKAAR	GLNVGHLKLT	FCSVGPTECA
78		ILR RPVALQLDY	N SVGDIGVEQI	LPCLGVCKAL	YLRDNNISDR	GICKLIECAL
84		LAL FNNKLTDGC	A HSMAKLLACF	QNFLALRLGN	NYITAAGAQV	LAEGLRGNTS
90		enr vgdegaqai	A EALGDHQSLF	R WLSLVGNNIG	SVGAQALALM	LAKNVMLEEL
96	CLEENHLO	QDE GVCSLAEG	LK KNSSLKILK	L SNNCITYLO	ga eallqa <u>p*</u>	
10	0.7					

# Figure 23 SEQ ID NO: 56

atqqqqqaaqaqqqtgqttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg aqaqtgtcctggactggctgctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct  $\verb|ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc|$ ccaaqaaqcccaqqccqacaqccaqtcccccaaqctqcatqqctqctqqqacccccactcqctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcqtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tgccacqqtqaaaqcqaatqqattqqctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagetgccacatgcaagaagtatatggccaagetgaggaccacggtgtctgctcagtctcgcttcctcagtacctat qatqqaqcaqaqacqctctgcctgqaggacatatacacaqagaatgtcctggaggtctgggcagatgtgggcatggctgg acccccqcaqaaqaqcccaqccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctqtqctqqtqqtqagqcqggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctctgaacagggcatcgagc tqtacctqaqqaaqcqccatcatqaqcccqqqqtqqcqqaccqcctcatccgcctgctccaagagacctcagccctgcac  $\verb|ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc|$ aaagaccactacagatatgtacctgctgattctgcagcatttttctgctgcatgccaccccccagactcagcttcccaag qtctqqqacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg tqctqctacqtqttctcaqcccagcagctccaqgcacacaggtcaqccctgatgacatttctcttggcttcctggtgcg tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg qccaqqctcctqcccacqatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga agacatetgagaaggeeetgetetggegeeaggeetgtgeeegetggtgtetggeeegeageeteegeaageaetteeae cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggccccgtggccctgcagctggactacaac tctgtgggtgacattggcgtggagcagctgcttgccttgccttggtgtctgcaaggctctgtattttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca a a ctacat cact g ccg cgg g ag ccca ag t g ctgg ccg ag g g ctccg ag g caa cacct cctt g cag t t cct g g g at t ctggggcaacagagtgggtgacgagggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcagtgtggggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtetggeteegagggaacaetttetetetagaggaggttgacaageteggetgeagggacaeeagaetettgett tga

## Figure 24 SEQ ID NO: 57

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGPPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLWRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

#### Figure 25 SEQ ID NO: 58

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga  $\verb|aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg|$ agagtgtcctggactggctgctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc ccaagaagcccaggccgacagccagtccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcqtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg acccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgggtgaggcgggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctqtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtgggcctgggcatg  ${\tt tgctgctacgtgttctcagcccagcagcaccaggcagcaccaggtcagccctgatgacatttctcttggcttcctggtgcg}$ tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga agacatctgagaaggccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tqqqccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac  $\verb|tctgtgggtgacattggcgtgagcagctgcttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat|$ ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca  ${\tt aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg}$  $\tt gcgcaacagagtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc$ tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc ${\tt tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat}$ cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgctt tga

# 1001 374 10001

SNP	Allele	SNP Sequence*	AA Polymorphism AA Location cDNA Location	AA Location	cDNA Location	
SNP4	-	GGCAGATGTGGCATGGCTGGACCC; SEQ ID NO:70	Д	P 268 S	nt 802	
	2	GGCAGATGTGGGCATGGCTGGAICC; SEQ ID NO:71	S			
SNP20	-	AGACATCTGAGAGGCCCTGCTCCGG; SEQ ID NO:72	R	R 702 W	nt 2104	
	2	AGACATCTGAGAAGGCCCTGCTCTGG; Seq ID NO:73	M			
SNP7	-	CTGCAGCACCTCCGGCGGCCCGTG: SEQ ID NO:74	Λ	V 793 M	nt 2377	
	2	CTGCAGCACCTCCGGCGGCCCATG; SEQ ID NO:75	M			
SNP18	1	TTGCAGAAGTTAGCTCTATTCAAC; SEQ ID NO:76	Z	N 852 S	nt 2555	
	2	TTGCAGAAGTTAGCTCTATTCAGC: SEQ ID NO:77	S			
SNP25	-	ACTGACGGCTGTGCACACTCCATG; SEQ ID NO:78	M	V 863 M	nt 2587	
	2	ACTGACGGCTGTGCACACTCCGTG; SEQ ID NO:79	Λ			
SNP17	1	TGCAGTTCCTGGGATTCTGGGGC; SEQ ID NO:80	9	G 908 R	nt 2722	
	2	TGCAGTTCCTGGGATTCTGGCGC; SEQ ID NO:81	R			
SNP23	1	CACTGATGCTGGCAAAGAACGTC; SEQ ID NO:82	Λ	V 955 I	nt 2863	
	2	CACTGATGCTGGCAAAGAACATC; SEQ ID NO:83	_			
3020C ins	1	GGGCAGAAGCCCTCCTGCAGGCCCT; SEQ ID NO:90	wild-type		nt 3020	
	2	GGGCAGAAGCCCTCCTGCAGGCCCCT; SEQ ID NO:91 frameshift mutation	frameshift mutation	ν33		
		*Underlined is mutated base				
		*Nucleotide/amino acid numbers designate the positions in Nod2a as reported by Ogura et al. J. Biol. Chem. 276:4812 [2001]	Nod2a as reported	by Ogura et al	J. Biol. Chem. 27	6:4812 [2001]
		AA = amino acid				
		SNP= single nucleotide polymorphism				

Figure 26

		snp23 3020insC comment	common	about 10% allele frequency in controls	about 4% allele frequency in controls	about 1-2% allele frequency in controls	about 3-4% allele frequency in controls	very rare: 4/400 patients are heterozygous; occurs on allele 1 of snp4	very rare: arms not working	very rare: arms working	very rare: arms working
		3020in	1	-	~	τ-	2	_	τ-	7	7
			1	-	-	_	-	-	-	-	0
		snp18 snp17	~	-	-	2	-	-	-	_	-
		snp18	-	-	1	-	-	-	-	7	-
		2dus	-	-	-	_	-	-	2	-	-
		snp20 snp7	-	-	2	-	-	-	-	-	_
		snp4	-	2	2	2	2	~	-	2	-

Figure 27

# Figure 28 SEQ ID NO: 59

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGPPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWRNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

## Figure 29 SEQ ID NO: 60

 $\verb|aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcaggggtccctggaaggcttcg|$  ${\tt agagtgtcctggactgctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct}$  $\verb|ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc|\\$ tg cagag teac cgg ceat tg teagg aggete cae agceat g tg gag ac at get gg acct gg cat gg gg gg teaching the second control of the settcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg atccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgggtgaggcgggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tqtacctgaqqaagcgccatcatgagccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg  ${\tt tgctgctacgtgttctcagcccagcagcaccaggcaccaggtcagccctgatgacatttctcttggcttcctggtgcg}$ tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga agacatetgagaaggeeetgeteeggegeeaggeetgtgeeegetggtgtetggeeegeageeteegeaageaetteeae tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggcacgtagcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac  $\verb|tctgtgggtgacattggcgtgagcagctgcttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat|$  ${\tt aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg}$ gggcaacagagtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc  ${\tt tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat}$  $\verb|cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc| \\$ tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgctt tga

# Figure 30 SEQ ID NO: 61

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGSPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

# Figure 31 SEQ ID NO: 62

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg agagtgtcctggactggctgctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttggggcctgtcagaagctcatcgcggctgc ccaagaagcccaggccgacagccagtccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg accccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgggtgaggcgggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg tgctgctacgtgttctcagcccagcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga agacatctgagaaggccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccatggccctgcagctggactacaac tctgtgggtgacattggcgtggagcagctgctgccttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg gggcaacagagtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgctt tga

## Figure 32 SEQ ID NO: 63

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGPPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPMALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

#### Figure 33 SEQ ID NO: 64

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg agagtgtcctggactggctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct  $\verb"ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc"$ ccaagaagcccaggccgacagccagtccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg accccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgaggcgggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtgggcctgggcatg tgctgctacgtgttctcagcccagcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg  ${\tt tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct}$ acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga agacatctgagaaggccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac tctgtgggtgacattggcgtggagcagctgctgccttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcagcaaca aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg gggcaacagagtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgctttga

# Figure 34 SEQ ID NO: 65

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGPPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFSNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

#### Figure 35 SEQ ID NO: 66

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga aatgtgetegeaggaggetttteaggeacagaggagceagetggtegagetgeteggteteagggtecetqqaaqgetteq agagtgtcctggactggctgctgtcctgggaggtcctctcctgggaggactacqagggcttccacctcctqqqccaqcct ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctqtcagaaqctcatcqcqqctqc ccaagaagcccaggccgacagccagtcccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaaqaaggctgcttgatct tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt  ${\tt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat}$ gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctgqaqqtctqqqcaqatqtqqqcatqqctqq acccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtggggtgaggcgggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcaqggcaa gacttccaggaatttctcttttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctqtqcqqac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccqcaaqqtqqtqaccaqccqtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaaqqqcttctctqaacaqqqcatcqaqc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaqqaactqttqctqcaqqaqqqqqqqtcccc aaagaccactacagatatgtacctgctgattctgcagcatttttctgctgcatqccacccccccaqactcaqcttcccaaq gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctggqcaqactqqctctqtqqqcctqqqcatq tgctgctacgtgttctcagcccagcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccqcgttct gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaqqccqa gccgcacaaccttcagatcacagcagccttcctggcagggctgttgtcccgggagcactggggcctqctqqctqaqtqcc agacatctg agaaggccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccactccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac  $\verb|ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca| \\$  ${\tt aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg}$ gggcaacagagtgggtgacgagggggcccaggccctggctgaagccttgggtgatcaccaqaqcttqaqqtqqctcaqcc  $\verb|tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacatcatgctagaagaactc|$ tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaaqccctcctqcaqqcccttqaaaqqaatqacaccatcc tggaagtetggeteegagggaacaetttetetetagaggaggttgacaageteggetgeagggacaceaqaetettqett tqa

# Figure 36 SEQ ID NO: 67

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGPPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNIMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

#### Figure 37 SEQ ID NO: 68

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg agagtgtcctggactggctgtcctgggagggtcctctcctgggaggactacgagggcttccacctcctgggccagcctctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc ccaagaagcccaggccgacagccagtcccccaagctgcatggctgctggggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagacatgctggacctggcatgggagcggggt ttcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct  ${\tt tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt}$ tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg acccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgggtgaggcggggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa  $\tt gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggactgcagtgcatggccaaaccactctctgtgcggactgcagtgcatggccaaaccactctctgtgcggactgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgcagtgcatggccaaaccactctctgtgcggactgcagtgc$ tetaetetttgageaetgetgttggeetgatgttggteaagaagaetetteeagttaeteettgaeeaceetgaeegtg acctetgtecagaccetgetetteaacettetgeagggeaacetgetgaagaatgeeegeaaggtggtgaeeageegtee ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtgggcctgggcatg tgctgctacgtgttctcagcccagcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg  ${\tt tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct}$ acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga gccgcacaacettcagatcacagcagcettcctggcagggctgttgtcccgggagcactggggcctqctqqctqaqtqcc agacatctg agaagg ccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac $\verb|cgagatgcaggagggggctggcatggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcag|\\$ tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctqqactacaa  $\verb|tctgtgggtgacattggcgtgagcagctgcttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat|$ ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg gggcaacagagtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc tgcctggaggagaaccatctccaggatgaaggtgtatgttctctccgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgctt

## Figure 38 SEQ ID NO: 69

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGPPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSVAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

## Figure 39 SEQ ID NO: 84

atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg aqaqtqtcctqqactqqctqctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc ccaagaagcccaggccgacagccagtcccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt  $\verb|ttcgtcagccagtatgaatgtgatatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct|$ tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg atccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggttgggtgaggcgggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac  $\verb|tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg|$ acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac qqtttqtqccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaaqaccactacagatatqtacctgctgattctgcagcattttctgctgcatgccaccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg tgctgctacgtgttctcagcccagcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg  ${\tt tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct}$ acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg  $\tt gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagctttgctgcagaaggccga$ agacatetgagaaggeeetgeteeggegeeaggeetgtgeeegetggtgtetggeeegeageeteegeaageaetteeae tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggagggggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg  ${\tt tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctgcactacaac}$ tctgtgggtgacattggcgtggagcagctgctgccttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg gcgcaacagaqtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcaggggacaccagactcttgctt tga

## Figure 40 SEQ ID NO: 85

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGSPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWRNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

## Figure 41 SEQ ID NO: 86

 $\verb|aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg| \\$ agagtgtcctggactgctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct $\verb|ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc|\\$ ccaagaagcccaggccgacagccagtcccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt ttcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt  ${\tt tggaagctgccacatgcaagaagtatatggccaagctgaagaccacggtgtctgctcagtctcgcttcctcagtacctat}$ gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg atccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtggggtgagggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgcatgccacccccccagactcagcttcccaag gtctgggacccagtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg tgctgctacgtgttctcagcccagcagctccaggcagcacaggtcagccctgatgacatttctcttggcttcctggtgcg tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga agacatctgagaaggccctgctccggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggcaggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac tctgtgggtgacattggcgtggagcagctgctgccttgccttggtgtctgcaaggctctgtatttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcagcaaca aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg gggcaacagagtgggtgacgaggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc  $\verb|tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc|$ tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtctggctccgagggaacactttctctctagaggaggttgacaagctcggctgcagggacaccagactcttgctt tga

# Figure 42 SEQ ID NO: 87

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGSPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLRRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFSNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL

## Figure 43 SEQ ID NO: 88

 ${ t atgggggaagagggtggttcagcctctcacgatgaggaggaaagagcaagtgtcctcctcggacattctccgggttgtga}$ aatgtgctcgcaggaggcttttcaggcacagaggagccagctggtcgagctgctggtctcagggtccctggaaggcttcg agagtgtcctggactggctgctgtcctgggaggtcctctcctgggaggactacgagggcttccacctcctgggccagcct ctctcccacttggccaggcgccttctggacaccgtctggaataagggtacttgggcctgtcagaagctcatcgcggctgc ccaagaagcccaggccgacagcccccaagctgcatggctgctgggacccccactcgctccacccagcccgagacc tgcagagtcaccggccagccattgtcaggaggctccacagccatgtggagaacatgctggacctggcatgggagcggggt  $\verb|tcgtcagccagtatgaatgtgatgaaatcaggttgccgatcttcacaccgtcccagagggcaagaaggctgcttgatct|$ tgccacggtgaaagcgaatggattggctgccttccttctacaacatgttcaggaattaccagtcccattggccctgcctt tggaagctgccacatgcaagaagtatatggccaagctgaggaccacggtgtctgctcagtctcgcttcctcagtacctat gatggagcagagacgctctgcctggaggacatatacacagagaatgtcctggaggtctgggcagatgtgggcatggctgg atccccgcagaagagcccagccaccctgggcctggaggagctcttcagcacccctggccacctcaatgacgatgcggaca ctgtgctggtggtgaggcgggcagtggcaagagcacgctcctgcagcggctgcacttgctgtgggctgcagggcaa gacttccaggaatttctctttgtcttcccattcagctgccggcagctgcagtgcatggccaaaccactctctgtgcggac tctactctttgagcactgctgttggcctgatgttggtcaagaagacatcttccagttactccttgaccaccctgaccgtg acctctgtccagaccctgctcttcaaccttctgcagggcaacctgctgaagaatgcccgcaaggtggtgaccagccgtcc ggccgctgtgtcggcgttcctcaggaagtacatccgcaccgagttcaacctcaagggcttctctgaacagggcatcgagc tgtacctgaggaagcgccatcatgagcccggggtggcggaccgcctcatccgcctgctccaagagacctcagccctgcac ggtttgtgccacctgcctgtcttctcatggatggtgtccaaatgccaccaggaactgttgctgcaggaggggggtcccc aaagaccactacagatatgtacctgctgattctgcagcattttctgctgcatgccaccccccagactcagcttcccaag qtctqqqacccaqtcttcttcggggccgcctccccaccctcctgcacctgggcagactggctctgtggggcctgggcatg  ${\tt tgctgctacgtgttctcagcccagcagcaccaggcaccaggtcagccctgatgacatttctcttggcttcctggtgcg}$  ${\tt tgccaaaggtgtcgtgccagggagtacggcgccctggaattccttcacatcactttccagtgcttctttgccgcgttct}$  ${\tt acctggcactcagtgctgatgtgccaccagctttgctcagacacctcttcaattgtggcaggccaggcaactcaccaatg}$  $\tt gccaggctcctgcccacgatgtgcatccaggcctcggagggaaaggacagcagcgtggcagctttgctgcagaaggccga$  $\tt gccgcacaaccttcagatcacagcagccttcctggcagggctgttgtcccgggagcactggggcctgctgagtgcc$ agacatctgagaaggccctgctctggcgccaggcctgtgcccgctggtgtctggcccgcagcctccgcaagcacttccac tccatcccgccagctgcaccgggtgaggccaagagcgtgcatgccatgcccgggttcatctggctcatccggagcctgta cgagatgcaggaggagcggctggctcggaaggctgcacgtggcctgaatgttgggcacctcaagttgacattttgcagtg tgggccccactgagtgtgctgccctggcctttgtgctgcagcacctccggcggcccgtggccctgcagctggactacaac tctgtgggtgacattggcgtggagcagctgctgccttgccttggtgtctgcaaggctctgtattttgcgcgataacaatat ctcagaccgaggcatctgcaagctcattgaatgtgctcttcactgcgagcaattgcagaagttagctctattcaacaaca  ${\tt aactacatcactgccgcgggagcccaagtgctggccgaggggctccgaggcaacacctccttgcagttcctgggattctg}$ gggcaacagagtgggtgacgagggggcccaggccctggctgaagccttgggtgatcaccagagcttgaggtggctcagcc tggtggggaacaacattggcagtgtgggtgcccaagccttggcactgatgctggcaaagaacgtcatgctagaagaactc tgcctggaggagaaccatctccaggatgaaggtgtatgttctctcgcagaaggactgaagaaaaattcaagtttgaaaat cctgaagttgtccaataactgcatcacctacctaggggcagaagccctcctgcaggcccttgaaaggaatgacaccatcc tggaagtetggeteegagggaacaetttetetetagaggaggttgacaageteggetgeagggacaceagaetettgett

#### Figure 44 SEQ ID NO: 89

MGEEGGSASHDEEERASVLLGHSPGCEMCSQEAFQAQRSQLVELLVSGSLEGFESVLDWLLSWEVLSWEDYEGFHLLGQP
LSHLARRLLDTVWNKGTWACQKLIAAAQEAQADSQSPKLHGCWDPHSLHPARDLQSHRPAIVRRLHSHVENMLDLAWERG
FVSQYECDEIRLPIFTPSQRARRLLDLATVKANGLAAFLLQHVQELPVPLALPLEAATCKKYMAKLRTTVSAQSRFLSTY
DGAETLCLEDIYTENVLEVWADVGMAGSPQKSPATLGLEELFSTPGHLNDDADTVLVVGEAGSGKSTLLQRLHLLWAAGQ
DFQEFLFVFPFSCRQLQCMAKPLSVRTLLFEHCCWPDVGQEDIFQLLLDHPDRVLLTFDGFDEFKFRFTDRERHCSPTDP
TSVQTLLFNLLQGNLLKNARKVVTSRPAAVSAFLRKYIRTEFNLKGFSEQGIELYLRKRHHEPGVADRLIRLLQETSALH
GLCHLPVFSWMVSKCHQELLLQEGGSPKTTTDMYLLILQHFLLHATPPDSASQGLGPSLLRGRLPTLLHLGRLALWGLGM
CCYVFSAQQLQAAQVSPDDISLGFLVRAKGVVPGSTAPLEFLHITFQCFFAAFYLALSADVPPALLRHLFNCGRPGNSPM
ARLLPTMCIQASEGKDSSVAALLQKAEPHNLQITAAFLAGLLSREHWGLLAECQTSEKALLWRQACARWCLARSLRKHFH
SIPPAAPGEAKSVHAMPGFIWLIRSLYEMQEERLARKAARGLNVGHLKLTFCSVGPTECAALAFVLQHLRRPVALQLDYN
SVGDIGVEQLLPCLGVCKALYLRDNNISDRGICKLIECALHCEQLQKLALFNNKLTDGCAHSMAKLLACRQNFLALRLGN
NYITAAGAQVLAEGLRGNTSLQFLGFWGNRVGDEGAQALAEALGDHQSLRWLSLVGNNIGSVGAQALALMLAKNVMLEEL
CLEENHLQDEGVCSLAEGLKKNSSLKILKLSNNCITYLGAEALLQALERNDTILEVWLRGNTFSLEEVDKLGCRDTRLLL